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<p>(21) International Application Number: PCT/IL98/00021</p> <p>(22) International Filing Date: 15 January 1998 (15.01.98)</p> <p>(30) Priority Data: 120022 16 January 1997 (16.01.97) IL</p> <p>(71) Applicants (for all designated States except US): YISSUM RESEARCH DEVELOPMENT COMPANY OF THE HEBREW UNIVERSITY OF JERUSALEM [IL/IL]; Jabotinsky Street 46, 91042 Jerusalem (IL). CONSEJO SUPERIOR DE INVESTIGACIONES CIENTIFICAS [ES/ES]; Calle Serrano, 117, E-28006 Madrid (ES).</p> <p>(72) Inventors; and (75) Inventors/Applicants (for US only): AVNIR, David [IL/IL]; Novomiesky Moshe Street 2, 96908 Jerusalem (IL). COHEN, Levy, David [ES/ES]; Santo Angel, 81, escalera 3, 1A, E-28043 Madrid (ES).</p> <p>(74) Agent: NOAM, Meir; P.O. Box 32081, 91320 Jerusalem (IL).</p>		<p>(81) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG).</p> <p>Published With international search report. Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.</p>
<p>(54) Title: SUNSCREENS FOR PROTECTION FROM SUN RADIATION</p> <p>(57) Abstract</p> <p>The present invention relates to sunscreen-doped sol-gel materials useful for protecting body tissues, such as skin, nails and hair and other surfaces from sunlight radiation. The sol-gel matrices are transparent to the UV radiation in the range above 250 nm and the doped sunscreen agents are either chemical or physical sunscreens capable of absorbing the UV radiation in the range above 250 nm. Any sunscreen molecule, moiety or particle may be used in the present invention. The sol-gel matrices may be particles in any shape, 0.01 - 100 microns in diameter, or they may be thin films, thin coatings or in the form of a monolith. The present invention also relates to a method for the preparation of sunscreen-doped sol-gel materials comprising condensation-polymerizing of at least one monomer selected from metal alkoxides, semi metal alkoxides, metal esters, semi metal esters and from monomers of the formula M(R)_n(P)_m, wherein M is a metallic or semi metallic element, R is a hydrolyzable substituent, n is an integer from 2 to 6, P is a non polymerizable substituent or a suncreening moiety or derivative and m is an integer from 0 to 6, in the presence of at least one sunscreen ingredient, resulting in the entrapment of the sunscreen ingredients within the formed sol-gel matrix.</p>		

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SUNSCREENS FOR PROTECTION FROM SUN RADIATION

Field of the Invention

The present invention generally relates to sunscreen - doped sol - gel materials useful for protecting body tissues and other surfaces from ultra violet radiation (the term "ultra violet radiation" in the present invention, unless otherwise specified, includes radiation in the range of 320-400nm, called UVA, and radiation in the range of 280-320 nm, called UVB. Ultra violet radiation is hereinafter called UV radiation), to a method for preparing said sunscreen - doped sol - gel materials and to a method for protecting body tissues and other surfaces from UV radiation, using the said sunscreens. More specifically, the present invention relates to sol - gel matrices produced by hydrolysis and condensation- polymerization of metal and semi-metal alkoxide monomers to which sunscreen molecules are added either prior to or after hydrolysis of the monomer, followed, if needed, by a pH change and by gelation. The resulting sunscreen - doped sol - gel materials can be obtained in almost any shape and may be applied, as dispersions or powders, to body tissues and other surfaces, thereby achieving protection of the body tissues and surfaces against damaging effects of the sun radiation, without having undesirable direct contact between the suncreening chemical ingredients and the body tissues.

Background of the Invention

Sunscreens are chemical ingredients which are usually applied topically to the skin and hair in order to protect them from the damaging effects of the sun's radiation, especially against (UV) ultraviolet radiation.

There is evidence that prolonged sunlight exposure results in increasing incidence of skin carcinogenesis, pigmentation, anomalies and precancerous lesions such as

actinic keratosis, melanoma and nonmelanoma skin cancers, as well as accelerated skin aging and undesirable changes in hair quality.

Sunscreens are usually classified in two major categories. The first includes chemical sunscreens which absorb sun radiation and therefore reduce the amount of UV radiation reaching the skin. Chemical sunscreens can be subdivided into eight derivative families: para amino benzoates, salicylates, cinnamates, benzophenones, anthranilates, dibenzoylmethanes, camphores and miscellaneous chemicals. The second category includes physical sunscreens which reflect, scatter or physically block the UV light reaching the skin surface. Physical sunscreens are mainly metal oxides such as titanium dioxide, zinc oxide and also red petrolatum.

Ultimate sunscreens should be chemically inert, highly photostable and neither sensitizing nor photosensitizing. Yet, according to more than 200 published reports relating to all major chemical sunscreen families, sunscreen agents induce both contact and photocontact dermatitis (Dromgoole and Maibach, In Sunscreens, eds. N.J. Lowe, N.A. Saath; Marcel Dekker: N.Y., 1990, Chapter 20). In other publications it was reported that commonly used sunscreen compounds undergo photodecomposition reactions (Roscher et al. J. Photochem. Photobiol. A : Chem, 80: 417-421, 1994). Moreover, although more people are using sunscreens, the rate of skin cancer around the world is rising. A possible explanation is that sunscreens might encourage, rather than prevent, sun related cancers. Indeed, it was found (Knowland et al. FEBS Letters, 324: 309-313, 1993) that the commercial sunscreen ingredient Padimate - O is mutagenic in sunlight. The sunlight - excited Padimate - O is thought to produce free radicals which directly attack the DNA.

Physical sunscreens, especially titania, although considered to be "relatively safe", are known to be photosensitizers capable of rupturing covalent bonds (U. Stafford, K.A. Gray and P.V. Kamat, Heterogeneous Chem Rev, 3, 77-104, 1996).

There is, therefore, a great need for isolating chemical and physical sunscreen agents from the body while retaining both the sunscreen activity and the compatibility with cosmetic preparations.

US Patent No. 5,223,250 and WO Patent 95/28912 describe the entrapment of sunscreen ingredients in organic polymers. However, organic polymers suffer a major drawback in that they do not provide photostability, a most important requirement of a sunscreen matrix. The photodegradation, photooxidation and photoreactivity of organic polymers are well documented (summarized in J.F. Rabeck, Photodegradation of polymers; Springer: Berlin, 1996). Also, plastic carriers, as in the above mentioned patents, are unable to tightly entrap dopants. Small molecules are known to diffuse in and out of these polymers, making the isolation from body tissues incomplete.

The present invention relates to sunscreen - doped sol - gel materials, such as chemical and physical sunscreens that are doped in a transparent sol-gel matrix, for the purpose of protecting body tissues (the term "body tissues" in the present invention refers to body tissues that are exposed to sun light such as hair, skin and nails) and other surfaces (the term "surfaces" in the present invention refers to surfaces that are exposed to sun light and may be damaged from the exposure to UV radiation) from ultra violet (UV) radiation, to a method for the preparation of these sunscreen - doped sol - gel materials and to a method for the protection of body tissues and other surfaces exposed to sunlight, from UV radiation, using the said sunscreens.

In the present invention the term "sunscreen - doped sol - gel" refers to sunscreen molecules or polymers (capable of absorbing or deflecting UV radiation) when they are doped in sol - gel matrix.

The term "chemical sunscreen" in the present invention refers to chemical ingredients which absorb sun radiation and therefore reduce the amount of UV radiation reaching the skin or other surfaces. The term "physical sunscreens" in

the present invention refers to chemical ingredients which reflect, scatter or physically block the UV light reaching the skin or other surfaces.

US patents numbers 5,292,801 and 5,300,564 describe the preparation of the inorganic porous ceramic glass - like sol - gel matrix obtained by hydrolysis and condensation - polymerization of metal and semi metal alkoxides resulting in oxide - type materials such as SiO_2 , TiO_2 , Al_2O_3 , ZnO and similar materials, as well as in their mixtures and their covalent bound organic derivatives. Due to the relatively low temperature needed for the preparation of sol - gel matrices, organic molecules can be doped into the matrix.

The sol-gel matrix of the present invention is thermally and photochemically stable, as opposed to the above mentioned plastic carriers, and can easily withstand normal duration of exposure to sunlight. Also, as opposed to the plastic carriers of the above patents, no leaching of the trapped sunscreen agents occurs from the sol-gel matrix, so that there is no direct contact of the sunscreen molecules with the body tissues.

The present invention relates to a simple method for preparing a matrix entrapping any sunscreen molecule or particle, providing, for the first time, a stable, safe and environment friendly sunscreen.

Summary of the Invention

The present invention relates to sunscreen-doped sol-gel materials useful for protecting body tissues, such as skin, nails and hair and other surfaces from sunlight radiation.

The sol-gel matrices are transparent to the UV radiation in the range above 250 nm and the doped sunscreen agents are either chemical or physical sunscreens capable of absorbing the UV radiation in the range above 250nm. Chemical sunscreens may be para amino benzoates, salicylates, cinnamates, benzophenones, anthranilates, dibenzoylmethanes, camphores and physical sunscreens may be titanium dioxide, zinc oxide and red petrolatum. This

invention is not limited in any way to the given examples of sunscreens agents. Any sunscreen molecule, moiety or particle may be used in the present invention. The sol-gel matrices may be particles in any shape, 0.01 - 100 microns in diameter, or they may be thin films, thin coatings or in the form of a monolith. The present invention also relates to a method for the preparation of sunscreen-doped sol-gel materials comprising condensation - polymerizing of at least one monomer selected from metal alkoxides, semi metal alkoxides, metal esters, semi metal esters and from monomers of the formula $M(R)_n(P)_m$, wherein M is a metallic or semi metallic element (such as silicon, titanium, zinc, aluminum, zirconium) R is a hydrolyzable substituent (such as alkoxides, aryloxides, carboxylic esters, acyloxy groups, diketonato groups, hydrolyzable aza groups and chlorine), n is an integer from 2 to 6, P is a non polymerizable substituent or a suncreening moiety or derivative and m is an integer from 0 to 6, in the presence of at least one sunscreen ingredient, resulting in the entrapment of the sunscreen ingredients within the formed sol - gel matrix.

Preferably, the sunscreen ingredients are added to the polymerizing mixture, either prior to or after hydrolysis of the monomer, followed, if needed, by a pH change and by gelation. The polymerization may performed also in the presence of a non - sunscreen functional co-dopant such as cosmetic ingredients or ingredients for coloring, leaching control, transparency/opacity control, acidity/basicity control, hydrophobicity/ hydrophilicity control.

The polymerization may be carried out under acidic, neutral or basic conditions for forming a porous gel, xerogel (dry gel) or colloidal suspension.

The present invention relates to a method for the protection of body tissues and other surfaces from UV radiation comprising coating said body tissues or surfaces with the sol - gel matrix entrapping sunscreen ingredients as described, and to cosmetic, pharmaceutical or industrial mixtures and compositions containing at least one type of doped sol - gel particle as described which is incorporated into a cosmetic or other vehicle to be applied topically to body tissues or to other surfaces.

Detailed Description of the Invention

The present invention relates to sunscreen - doped sol - gel materials useful for protecting body tissues and other surfaces from ultra violet (UV) radiation, to a method for preparing said sunscreens and to a method for protecting body tissues and other surfaces from UV radiation, using the said sunscreens. In the present invention sol - gel matrices are generated by hydrolysis and condensation-polymerization of metal and semi-metal alkoxide monomers to which sunscreen molecules are added either prior to or after hydrolysis of the monomer, followed, if needed, by a pH change and by gelation. The resulting sunscreen - doped sol - gel materials can be obtained in almost any shape and may be applied, as dispersions or powders, to body tissues or other surfaces, thereby achieving protection of the body tissues or surface against damaging effects of the sun radiation, without having undesirable direct contact between the suncreening chemical ingredients and the body tissues.

As opposed to any of the sunscreens known today, the sunscreen - doped sol-gel matrices of the present invention have the following advantages:

The sol - gel matrix is transparent to the UV range above 250nm, thus allowing the UVA and UVB light to penetrate the sol - gel particle and interact with the entrapped sunscreen molecule.

The sol - gel matrix is thermally and photochemically stable (in contrast to plastic carriers), easily withstanding the normal duration of exposure to sunlight and much beyond it.

The preparation of sunscreen - doped sol - gel matrices is simple; direct physical entrapment in the course of the sol - gel polymerization is possible and no reaction with the sunscreen molecule itself is needed.

The method of the present invention is not limited to the type of sunscreen ingredients used because any sunscreen molecule can be entrapped.

Points which are important for cosmetic preparations:

The sol - gel matrix can remain visibly transparent when applied on human tissues, as, for instance, in the case of SiO_2 sol - gel matrix.

Sunscreen - doped sol - gel particles can be obtained in various sizes, including downsizing to sub - micron levels. The particle can be prepared in various shapes, including spherical particles which are preferable for smooth contact with the skin. Practically any desired shape and form, from microscopic to macroscopic, is attainable.

The hydrophobicity/hydrophilicity ratio of the sol - gel particles surface can be controlled by the suitable choice of the monomers (e.g., using trialkoxy silane derivatives). Likewise, the acidity/basicity of the particles can be controlled.

The entrapped sunscreen molecules are not in direct contact with the skin. Each doped sunscreen molecule or particle is isolated in an individual cage, thus avoiding ground state or photochemical interactions with impurities, photodecomposition products, adjacent sunscreen molecules or other ingredients of the cosmetic vehicle. As a consequence, a sunscreen cosmetic preparation with a broad absorption spectrum can be obtained by the incorporation of at least two types of sunscreen - doped sol - gel particles, either by co - entrapment or by entrapment in different particles, which differ in their UV spectra. Since the doped sunscreen molecules have improved thermal and photochemical stability on exposure to sunlight, the shelf life of these materials and their cosmetic preparations or mixtures for other use, increases.

No leaching occurs of the trapped sunscreen agent from the matrix into the cosmetic vehicle.

The sunscreen - doped sol - gel matrix may have a broad absorption spectrum, encompassing both the UVA and UVB regions, if the doped sunscreens differ in their UV spectra. Due to their complete matrix isolation the entrapped sunscreens do not color or discolor skin and do not stain clothes. Also, the entrapped sunscreens are odorless when applied on skin or hair.

No deposition of sunscreen crystals is possible. Crystallization is a major cause for the need to use high sunscreen concentrations for desired sun protection factor (SPF) values.

The small particles of the doped sol - gel material blend optimally in cosmetic preparations.

Sol - gel matrices, such as silica, are non - allergenic and biocompatible. When the sunscreen cosmetic vehicle is washed off the body, the entrapped sunscreen is markedly more environmentally friendly than the free molecule, which becomes a pollutant. Silica is safe to the environment and so is the molecule entrapped within it.

The sol - gel matrix is basically obtained by condensation - polymerizing of monomers of which at least one monomer is selected from the group consisting of metal alkoxides, semi metal alkoxides, metal esters, semi metal esters and from monomers of the formula $M(R)_n(P)_m$, wherein M is a metallic or semi metallic element, R is a hydrolyzable substituent, n is an integer from 2 to 6, P is a non polymerized substituent and m is an integer from 0 to 6. The metallic or semi metallic element may be silicon, titanium, zinc, aluminum or zirconium, the hydrolyzable substituent may be from the group of alkoxides, aryloxides, carboxylic esters, acyloxy groups, diketonato groups, hydrolyzable aza groups and chlorine and the non polymerized substituent may be used for coloring, leaching control, transparency/opacity control, acidity/basicity control or hydrophobicity/hydrophilicity control or may be a sunscreen moiety or derivative.

The non - sunscreen co-dopants may be cosmetic ingredients or for coloring, leaching control, transparency/opacity control, acidity/basicity control, hydrophobicity/hydrophilicity control.

Examples of suitable monomers are for example, the alkoxysilanes (such as tetramethoxysilane, tetraethoxysilane, alkyl - trimethoxysilane etc.) resulting in silica - type (SiO_2) sol - gel glasses. Other preferred monomers are those which lead to the formation of ZnO and TiO_2 sol - gel matrices (e.g. zinc di - t - butoxide and titanium triisopropoxide).

The polycondensation is associated with gelation of a sol, which after drying is densified by a mild heat to form a solid matrix called xerogel. The optimal time to add the sunscreen molecules is either prior to or after hydrolysis of the monomer, followed, if needed, by a pH change, which is then followed by gelation. The properties of the final solid matrix are determined by the chemical and physical conditions during the preparation process, e.g. the ratio of metal/water (e.g. silane/ water), the pH, the absence or presence of catalysts, the temperature, the drying time and the incorporation of organic additives such as surface active agents. The doped material can be obtained in almost any shape, including spherical particles which are preferable for smooth contact with the skin.

Methods for obtaining fine dispersions and powders include: grinding; preventing the sol from gelation by dilution; pH changes; addition of surfactants; spraying techniques; and controlled growth, as is taught in the general field of colloids (L.H. Clint et al., Faraday Discuss. 95, 219, 1993). Multilayered particles are likewise obtainable.

The said invention will be further illustrated by the following experiments and examples. These experiments and examples do not intend to limit the scope of the invention but to demonstrate and clarify it only.

In a typical procedure, mixtures containing the alkoxysilane monomer (typically tetraethoxysilane or tetramethoxysilane (TMOS)) and water, are prepared at different ratios of silane/ water/ alcohol (such as ethanol or methanol). Examples for such mixtures are those containing 18 mmol of TMOS, 50 mmol of methanol and either 18 mmol or 72 mmol of distilled water. the mixtures can be made to polymerize with either basic or acidic catalysts. In a typical case, acid catalysis is induced by adding 0.425 mmol of HCl. At the sol stage, 0.10 to 0.64 mmol of the sunscreen are added, depending on the desired SPF (sunscreen materials are usually at concentrations suitable for SPF values between 2 and 50), any desired sunscreen can be added at this stage, with typical examples being para -

aminobenzoic acid (PABA) as a representative of the para amino benzoate sunscreens and octylmethoxycinnamate as an example of the cinnamate sunscreens. Other sunscreens usable at this stage are the salicylates, the benzophenones family, anthranilates, dibenzoylmethanes, camphores, metallic complexes and metal oxides.

Gelation is carried out at room temperature in a glass bottle covered with aluminum foil. After gelation takes place the aluminum foil is perforated, allowing slow evaporation of the solvent until dried xerogels are formed. The dried material is then ground to a fine powder. The doped xerogel microparticles can be coated with a second layer of sol - gel materials by dispersing in a mixture of TMOS/water/methanol with molar ratios 1/4/2.8, respectively, and stirring for six hours. The particles are separated and dried under vacuum for 10 hours at a temperature of 353K.

Leaching tests were performed by taking samples of 0.1 - 0.15 gr of each type of the doped particles and immersing in a flask containing 4 ml of methanol. The flask was covered and stirred for two hours at room temperature. The immersed particles were separated and dried for 10 hours at 323K. The entrapment proved to be leach proof and photostable, following prolonged exposure to sunlight (typically 7 hours).

Claims

- 1) Sunscreen-doped sol-gel materials useful for protecting body tissues and other surfaces from sunlight radiation.
- 2) Sunscreen-doped sol-gel materials according to claim 1 wherein the body tissues are skin, nails and hair.
- 3) Sunscreen-doped sol-gel materials according to claim 1 wherein the doped sunscreen materials are chemical sunscreen particles or molecules or physical sunscreens particles or molecules.
- 4) Sunscreen-doped sol-gel materials according to claim 3 wherein the doped sunscreen materials are capable of absorbing the UV radiation in the range above 250nm.
- 5) Sunscreen-doped sol-gel materials according to claim 3 wherein the chemical sunscreens are selected from para amino benzoates, salicylates, cinnamates, benzophenones, anthranilates, dibenzoylmethanes, camphores.
- 6) Sunscreen-doped sol-gel materials according to claim 3 wherein the physical sunscreens are UV reflecting or UV scattering entities.
- 7) Sunscreen-doped sol-gel materials according to claim 6 wherein the physical sunscreens are selected from titanium dioxide, zinc oxide and red petrolatum.
- 8) Sunscreen-doped sol-gel materials according to claim 1 wherein the sol-gel matrices are transparent to the UV radiation in the range above 250 nm, covering the UVA and UVB regions.

- 9) Sunscreen-doped sol-gel materials according to claim 1 wherein the sol-gel matrices are particles, either dried or dispersed as a sol.
- 10) Sunscreen-doped sol-gel materials according to claim 9 wherein the particles are 0.01 - 100 microns in diameter.
- 11) Sunscreen-doped sol-gel materials according to claim 9 wherein the sol-gel particles are spherically shaped.
- 12) Sunscreen-doped sol-gel materials according to claim 1 wherein the sol-gel matrices are thin films or thin coatings.
- 13) Sunscreen-doped sol-gel materials according to claim 1 wherein the sol-gel matrices are in the form of a monolith.
- 14) A method for the preparation of sunscreen-doped sol-gel materials comprising condensation - polymerizing of at least one monomer selected from metal alkoxides, semi metal alkoxides, metal esters, semi metal esters and from monomers of the formula $M(R)_n(P)_m$, wherein M is a metallic or semi metallic element, R is a hydrolyzable substituent, n is an integer from 2 to 6, P is a non polymerizable substituent or a suncreening moiety or derivative and m is an integer from 0 to 6, in the presence of at least one sunscreen ingredient, resulting in the entrapment of the sunscreen ingredients within the formed sol - gel matrix.
- 15) A method according to claim 14 wherein the sunscreen ingredients are added after hydrolysis of the monomer, followed by a pH change and gelation.
- 16) A method according to claim 14 wherein the sunscreen ingredients are added to the polymerizing mixture prior to hydrolysis of the monomer.

- 17) A method according to claim 14 wherein the polymerization is performed also in the presence of a non - sunscreen functional co-dopant.
- 18) A method according to claim 14 wherein the polymerization is carried out under acidic, neutral or basic conditions for forming a porous gel, xerogel (dry gel) or colloidal suspension.
- 19) A method according to claim 14 wherein the metallic or semi metallic element is selected from silicon, titanium, zinc, aluminum, zirconium and the hydrolyzable substituent is selected from alkoxides, aryloxides, carboxylic esters, acyloxy groups, diketonato groups, hydrolizable aza groups and chlorine and the non polymerizable substituent is for coloring, leaching control, transparency/opacity control, acidity/basicity control or hydrophobicity/hydrophilicity control or is a sunscreen moiety or derivative
- 20) A method according to claim 14 wherein the sunscreen ingredients are chemical sunscreens or physical sunscreens.
- 21) A method according to claim 20 wherein the chemical sunscreens are selected from para amino benzoates, salicylates, cinnamates, benzophenones, anthranilates, dibenzoylmethanes, camphores and miscellaneous chemicals .
- 22) A method according to claim 20 wherein the physical sunscreens are UV reflecting or UV scattering entities.
- 23) A method according to claim 20 wherein the physical sunscreen are selected from titanium dioxide, zinc oxide and red petrolatum.

- 24) A method according to claim 17 wherein the non - sunscreen co-dopants are cosmetic ingredients or for coloring, leaching control, transparency/opacity control, acidity/basicity control, hydrophobicity/ hydrophilicity control.
- 25) A method according to claim 24 wherein the cosmetic ingredients are perfumes.
- 26) A method according to claim 14 wherein the sol - gel is obtained as a fine dispersion or as a powder by grinding or by stopping the sol from gelation by dilution or by pH changes or by adding surfactants or by using spraying techniques or by using controlled growth.
- 27) A method according to claim 14 wherein the sunscreen materials are at concentrations suitable for SPF values between 2 and 50.
- 28) A method for the protection of body tissues and other surfaces from UV radiation comprising coating said body tissues or surfaces with the sol - gel matrix entrapping sunscreen ingredients according to claim 1.
- 29) Cosmetic , pharmaceutical or industrial mixtures and compositions containing at least one type of doped particle according to claim 1 which is incorporated into a cosmetic or other vehicle to be applied topically to body tissues or to other surfaces.
- 30) Sunscreen-doped sol-gel materials as hereinbefore described.
- 31) A method for the preparation of sunscreen-doped sol-gel materials as hereinbefore described

INTERNATIONAL SEARCH REPORT

International Application No

PCT/IL 98/00021

A. CLASSIFICATION OF SUBJECT MATTER

IPC 6 A61K7/42

According to International Patent Classification(IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 6 A61K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	EP 0 281 034 A (T.YAMAMOTO C/O NAKATO LABORATORY) 7 September 1988 see claims 1,4,6,7 see page 6, line 38-41 ---	1,2,9, 14, 16-19, 24,25, 29,31
X	PATENT ABSTRACTS OF JAPAN vol. 096, no. 008, 30 August 1996 & JP 08 099838 A (NIPPON SHIKIZAI KOGYO KENKYUSHO), 16 April 1996, see abstract --- -/--	1-3,14, 16-20, 24,29-31

☒ Further documents are listed in the continuation of box C.

☒ Patent family members are listed in annex.

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INTERNATIONAL SEARCH REPORT

Intern 1al Application No

PCT/IL 98/00021

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	CHEMICAL ABSTRACTS, vol. 124, no. 12, 18 March 1996 Columbus, Ohio, US; abstract no. 153660, K.DOSHITA E.A.: "Glass flakes containing dispersed metal oxides, especially for use in cosmetics and sol-gel process for their manufacture" XP002065558 see abstract	1-3,6,7, 9,14,16, 18-20, 22,23, 28-31
X	& JP 07 315 859 A (NIPPON SHEET GLASS) -----	
X	CHEMICAL ABSTRACTS, vol. 119, no. 26, 27 December 1993 Columbus, Ohio, US; abstract no. 273503, K.MIZOHATA: "granular coloring materialsof organic-inorganic composites" XP002065559 see abstract	1-3,9, 14, 16-19,24
X	& JP 05 178 995 A (NIPPON PAINT) -----	

INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No

PCT/IL 98/00021

Patent document cited in search report		Publication date	Patent family member(s)	Publication date
EP 281034	A	07-09-1988	JP 2566386 B	25-12-1996
			JP 7268383 A	17-10-1995
			JP 2588236 B	05-03-1997
			JP 64000161 A	05-01-1989
			US 4880851 A	14-11-1989
			US 4988744 A	29-01-1991
			US 4980392 A	25-12-1990
			US 4987161 A	22-01-1991
			US 5075350 A	24-12-1991
			US 5387622 A	07-02-1995
			JP 2619905 B	11-06-1997
			JP 64000126 A	05-01-1989
			DE 3854796 D	01-02-1996
			DE 3854796 T	02-05-1996
			EP 0287877 A	26-10-1988
			JP 1026640 A	27-01-1989
			JP 2619910 B	11-06-1997
			US 4889876 A	26-12-1989